

Syngpl20mn

1 CTCGAGATCC ATTGTGCTCT AAAGGAGATA CCCGGCCAGA CACCCTCACC
51 TGCGGTGCCC AGCTGCCCAG GCTGAGGCAA GAGAAGGCCA GAAACCATGC
101 CCATGGGGTC TGTGCAACCG CTGGCCACCT TGTACCTGCT GGGGATGCTG
151 GTCGCTTCCG TGCTAGCCAC CGAGAAGCTG TGGGTGACCG TGTACTACGG
201 CGTGCCCGTG TGAAGGAGG CCACCACCAC CCTGTTCTGC GCCAGCGACG
251 CCAAGGCGTA CACACCGAG GTGCACAACG TGTGGGCCAC CCAGGCGTGC
301 GTGCCCACCG ACCCCAACCC CCAGGAGGTG GAGCTCGTGA ACGTGACCGA
351 GAACTTCAAC ATGTGGAAGA ACAACATGGT GGAGCAGATG CATGAGGACA
401 TCATCAGCCT GTGGGACCAG AGCCTGAAGC CCTGCGTGAA GCTGACCCCC
451 CTGTGCGTGA CCTGAACTG CACCGACCTG AGGAACACCA CCAACACCAA
501 CAACAGCACC GCCAACAACA ACAGCAACAG CGAGGGCACC ATCAAGGGCG
551 GCGAGATGAA CAACTGCAGC TTCAACATCA CCACCAGCAT CCGCGACAAG
601 ATGCAGAAGG AGTACGCCCT GCTGTACAAG CTGGATATCG TGAGCATCGA
651 CAACGACAGC ACCAGCTACC GCCTGATCTC CTGCAACACC AGCGTGATCA
701 CCCAGGCCTG QCCCAAGATC AGCTTCGAGC CCATCCCCAT CCACTACTGC
751 GCCCCCGCCG GCTTCGCCAT CCTGAAGTGC AACGACAAGA AGTTCAGCGG
801 CAAGGGCAGC TGCAAGAACG TGAGCACCGT GCAGTGCACC CACGGCATCC
851 GGCCGGTGGT GAGCACCCAG CTCCTGCTGA ACGGCAGCCT GGCCGAGGAG
901 GAGGTGGTGA TCCGCAGCGA GAACTTCACC GACAACGCCA AGACCATCAT
951 CGTGACACCTG AATGAGAGCG TGCAGATCAA CTGCACGCGT CCCAACTACA
1001 ACAAGCGCAA GCGCATCCAC ATCGGCCCCG GCGCGCCTT CTACACCACC
1051 AAGAACATCA TCGGCACCAT CCGCCAGGCC CACTGCAACA TCTCTAGAGC
1101 CAAGTGAAC GACACCCTGC GCCAGATCGT GAGCAAGCTG AAGGAGCAGT
1151 TCAAGAACAA GACCATCGTG TTCAACCAGA GCAGCGGCGG CGACCCCGAG
1201 ATCGTGATGC ACAGCTTCAA CTGCGGCGGC GAATTCTTCT ACTGCAACAC
1251 CAGCCCCCTG TTCAACAGCA CCTGGAACGG CAACAACACC TGAACAACA
1301 CCACCGGCAG CAACAACAAT ATTACCCTCC AGTGCAAGAT CAAGCAGATC
1351 ATCAACATGT GGCAGGAGGT GGGCAAGGCC ATGTACGCCC CCCCCATCGA
1401 GGGCCAGATC CGGTGCAGCA GCAACATCAC CGGTCTGCTG CTGACCCGCG
1451 ACGGCGGCAA GGACACCGAC ACCAACGACA CCGAAATCTT CCGCCCCGGC

FIG 1
(SHEET 1 OF 4)

1501 GGGGGCGACA TCGCGGACAA CTGGAGATCT GAGCTGTACA AGTACAAGGT
1551 GGTGACGATC GAGCCCCTGG GCGTGGCCCC CACCAAGGCC AAGCGCCGCG
1601 TGGTGCAGCG CGAGAAGCGC TAAAGCGGCC GC (SEQ ID NO:34)

0871234.09096

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1 ACCGAGAAGC TGTGGGTGAC CGTGTACTAC GCGGTGCCCG TGTGGAAGGA
51 GGCCACCACC ACCCTGTTCT GCGCCAGCGA CGCCAAGGCG TACGACACCG
101 AGGTGCACAA GGTGTGGGCC ACCCAGGCGT GCGTGCCAC CGACCCCAAC
151 CCGCAGGAGG TGGAGCTCGT GAACGTGACC GAGAACTTCA ACATGTGGAA
201 GAACAACATG CTGGAGCAGA TGCATGAGGA CATCATCAGC CTGTGGGACC
251 AGAGCCTGAA GCGCTGCGTG AAGCTGACCC CCCTGTGCGT GACCCCTCAAC
301 TGCACCGACC TGAGGAACAC CACCAACACC AACAACAGCA CCGCCACAA
351 CAACAGCAAC AGCGAGGGCA CCATCAAGGG CGGCGAGATG AAGAACTGCA
401 GCTTCAACAT CACCACCAGC ATCCGCGACA AGATCCAGAA GGAGTACGCC
451 CTGCTGTACA AGCTGGATAT CGTGAGCATC CACAACGACA GCACCAGCTA
501 CCGCCTGATC TCCTGCAACA CCAGCGTGAT CACCCAGGCC TCGCCCAAGA
551 TCAGCTTCGA GCGCATCCCC ATCCACTACT GCGCCCCCGC CGGCTTCGCC
601 ATCCTGAAGT GCAACGACAA GAAGTTCAGC GGCAAGGGCA GCTGCAAGAA
651 CGTGACCACC GTGCAGTGCA CCCACGGCAT CCGGCCGGTG GTGAGCACCC
701 ACCTCCTGCT GAACGGCAGC CTGCGCGAGG AGGAGGTGGT GATCGGCAGC
751 GAGAACTTCA CCGACAACGC CAAGACCATC ATCGTGACCC TGAATGAGAG
801 CGTGCAGATC AACTGCACGC GTCCCAACTA CAACAAGCGC AAGCGCATCC
851 ACATCGGCCC CGGGCGCGCC TTCTACACCA CCAAGAACAT CATCGGCACC
901 ATCCGCGCAGG CCGACTGCAA CATCTCTAGA GCCAAGTGGA ACGACACCCT
951 GCGCCAGATC GTGAGCAAGC TGAAGGAGCA GTTCAAGAAC AAGACCATCG
1001 TGTTCACCA GAGCAGCGGC GCGGACCCCG AGATCGTGAT GCACAGCTTC
1051 AACTGCGGCG GCGAATTCTT CTACTGCAAC ACCAGCCCCC TGTTC AACAG
1101 CACCTGGAAC GGCAACAACA CCTGGAACAA CACCACCGGC AGCAACAACA
1151 ATATTACCCT CCAGTGCAAG ATCAAGCAGA TCATCAACAT GTGGCAGGAG
1201 GTGGGCAAGG CCATGTACGC CCCCCCATC GAGGGCCAGA TCCGGTGCAG
1251 CAGCAACATC ACCCGTCTGC TGCTGACCCG CGACGGCGGC AAGGACACCG
1301 ACACCAACGA CACCGAAATC TTCCGCCCCG GCGGCGGCGA CATGCGCGAC
1351 AACTGGAGAT CTGAGCTGTA CAAGTACAAG GTGGTGACGA TCGAGCCCCT
1401 GGGCGTGGCC CCCACEAAGG CCAAGCGCCG CGTGGTGACG CCGGAGAAGC

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1451 GGGCCGCCAT CCGCGCCCTG TTCCTGGGCT TCCTGGGGGC GGCGGGCAGC
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1551 GAGCGGCATC GTGCAGCAGC AGAACAACCT CCTCCGCGCC ATCGAGGCCC
1601 AGCAGCATAT GCTCCAGCTC ACCGTGTGGG GCATCAAGCA GCTCCAGGCC
1651 CGCGTGCTGG CCGTGGAGCG CTACCTGAAG GACCAGCAGC TCCTGGGCTT
1701 CTGGGGCTGC TCCGGCAAGC TGATCTGCAC CACACCGGTA CCCTGGAACG
1751 CCTCCTGGAG CAACAAGAGC CTGGACGACA TCTGGAACAA CATGACCTGG
1801 ATGCAGTGGG ACGCGAGAT CGATAACTAC ACCAGCCTGA TCTACAGCCT
1851 GCTGGAGAAG ACCCAGACCC AGCAGGAGAA GAAAGAGCAG GAGCTGCTGG
1901 AGCTGGACAA CTGGGCGAGC CTGTGGAAC TGTTCGACAT CACCAACTGG
1951 CTGTGGTACA TCAAAATCTT CATCATGATT GTGGGCGGCC TGGTGGGCCT
2001 CCGCATCGTG TTCGCCGTGC TGAGCATCGT GAACCGCGTG CGCCAGGGCT
2051 ACAGCCCCCT GAGCCTCCAG ACCCGGCCCC CCGTGCCGCG CGGGCCCGAC
2101 CGCCCCGAGG GCATCGAGGA GGAGGGCGGC GAGCGCGACC GCGACACCAG
2151 CGGCAGGCTC GTGCACGGCT TCCTGGCGAT CATCTGGGTC GACCTCCGCA
2201 GCCTGTTCCT GTTCAGCTAC CACCACCGCG ACCTGCTGCT GATCGCCGCC
2251 CGCATCGTGG AACTCCTAGG CCGCCGCGGC TGGGAGGTGC TGAAGTACTG
2301 GTGGAACCTC CTCCAGTATT GGAGCCAGGA GCTGAAGTCC AGCGCCGTGA
2351 GCCTGCTGAA CGCCACCGCC ATCGCCGTGG CCGAGGGCAC CGACCGCGTG
2401 ATCGAGGTGC TCCAGAGGGC CGGGAGGGCG ATCCTGCACA TCCCCACCCG
2451 CATCCGCCAG CGGCTCGAGA GGGCGCTGCT G (SEQ ID NO:35)

001717294.000000

FIG. 1

(SHEET 4 OF 4)

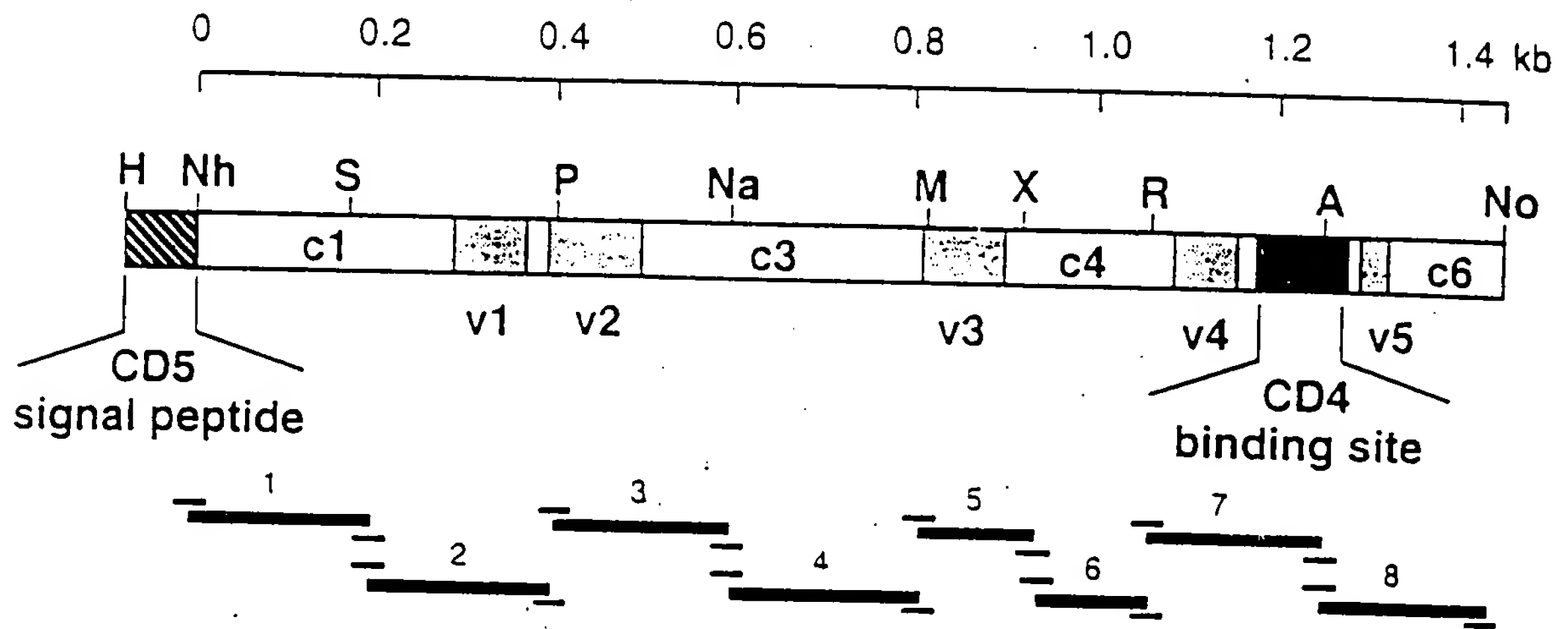


FIGURE 2

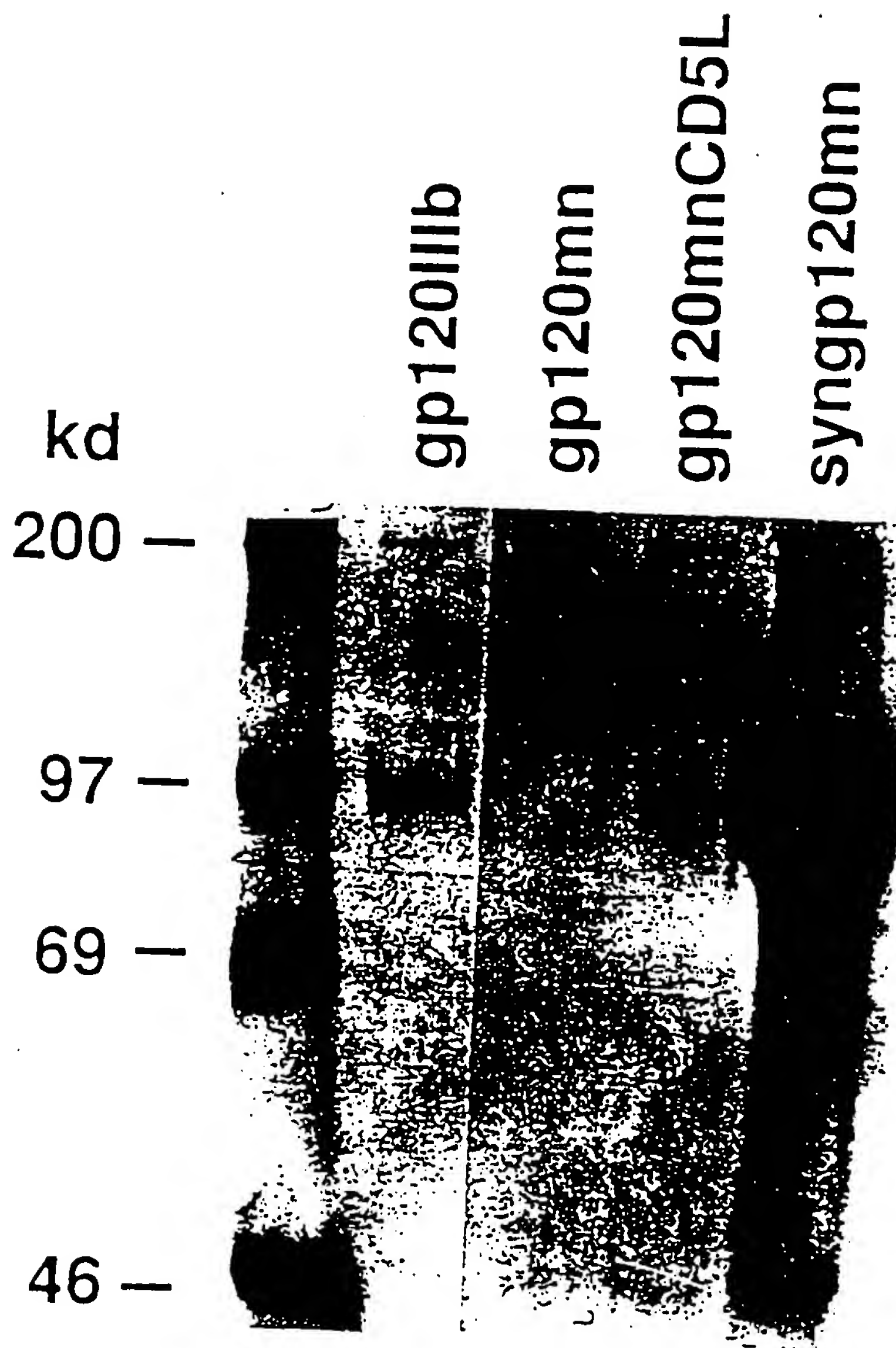


FIGURE 3

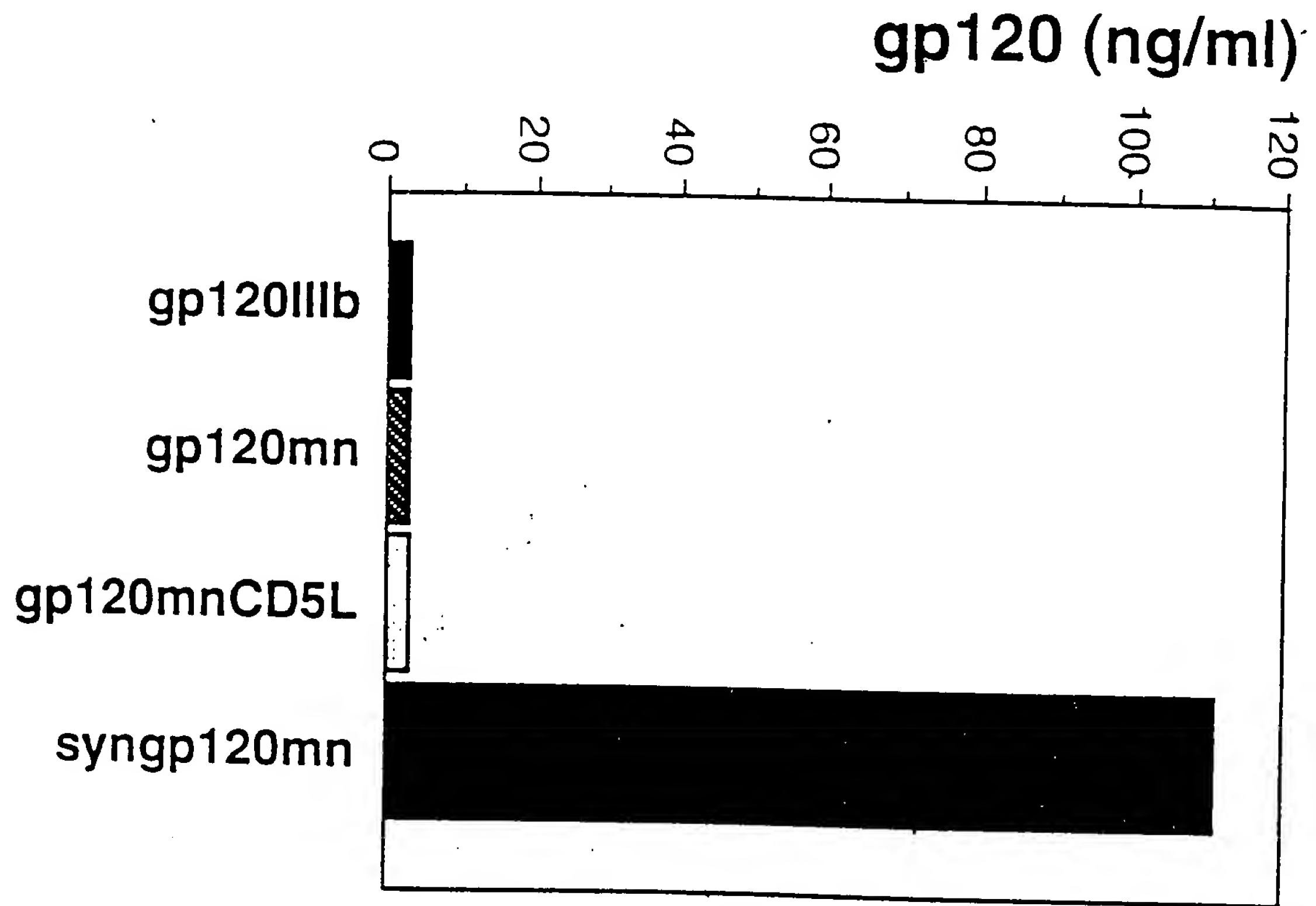


FIGURE 4

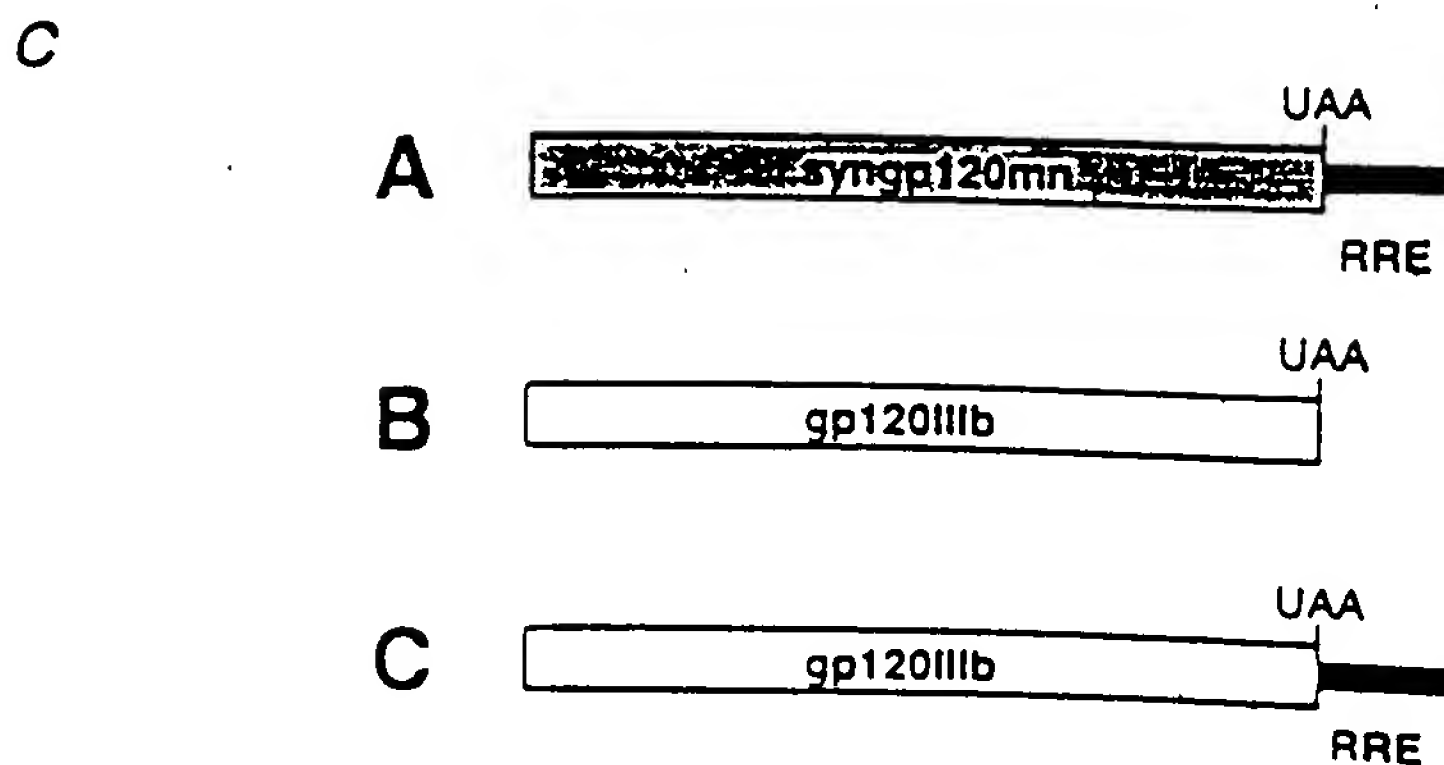


FIGURE 5

00717294-000000

(SEQ ID NO:36) env	→	atg	aat	cca	gta	ata	ata	agt	ata	aca	tta	tta	tta	agt	gta	tta	caa	atg	agt	aga	gga	caa
(SEQ ID NO:37) wt	→	atg	aac	cca	gtc	atc	atc	agc	atc	act	ctc	ctg	ctt	tca	gtc	tbg	cag	atg	tcc	cga	gga	cag
		R	V	I	S	L	T	A	C	L	V	N	Q	N	L	R	L	D	C	R	H	
env		aga	gta	ata	agt	tta	aca	gca	tgt	tta	gta	aat	caa	aat	tbg	aga	tta	gat	tgt	aga	cat	
wt		agg	gtg	atc	agc	ctg	aca	gcc	tgc	ctg	gtg	aa	cag	aac	ctt	cga	ctg	gac	tgc	cgt	cat	
		E	N	N	T	N	L	P	I	Q	H	E	F	S	L	T	R	E	K	K	K	
env		gaa	aat	aat	aca	cct	tbg	cca	ata	caa	cat	gaa	ttt	tca	tta	acg	cgt	gaa	aaa	aaa	aaa	
wt		gag	aat	aac	acc	aac	tbg	ccc	atc	cag	cat	gag	tbc	agc	ctg	acc	cga	gag	aag	aag	aag	
		H	V	L	S	G	T	L	G	V	P	E	H	T	Y	R	S	R	V	N	L	
env		cat	gta	tta	agt	gga	aca	tta	gga	gta	cca	gaa	cat	aca	tat	aga	agt	aga	gta	aat	tbg	
wt		cac	gtg	ctg	tca	ggc	acc	ctg	ggg	gtt	ccc	gag	cac	act	tac	cgc	tcc	cgc	gtc	aac	ctt	
		F	S	D	R	F	I	K	V	L	T	L	A	N	F	T	T	K	D	E	G	
env		ttt	agt	gat	aga	ttc	ata	aaa	gta	tta	aca	tta	gca	aat	ttt	aca	aca	aaa	gat	gaa	gga	
wt		ttc	agt	gac	cgc	ttt	atc	aag	gtc	ctt	act	cta	gcc	aac	ttc	acc	acc	aag	gat	gag	ggc	
		D	Y	M	C	E	L	R	V	S	G	Q	N	P	T	S	S	N	K	T	I	
env		gat	tat	atg	tgt	gag	ctc	aga	gta	agt	gga	caa	aat	cca	aca	agt	agt	aat	aaa	aca	ata	
wt		gac	tac	atg	tgt	gaa	ctt	cga	gtc	ctg	ggc	cag	aat	ccc	aca	agc	tcc	aat	aaa	act	atc	
		N	V	I	R	D	K	L	V	K	C	G	G	I	S	L	L	V	Q	N	T	
env		aat	gta	ata	aga	gat	aaa	tta	gta	aaa	tgt	gga	gga	ata	agt	tta	tta	gta	caa	aat	aca	
wt		aat	gtg	atc	aga	gac	aag	ctg	gtc	gtc	tgt	ggt	ggc	ata	agc	ctg	ctg	gtt	caa	aac	act	
		S	W	L	L	L	L	L	L	S	L	S	F	L	Q	A	T	D	F	I	S	
env		agt	tgg	tta	tta	tta	tta	tta	tta	agt	tta	agt	ttt	tta	caa	gca	aca	gat	ttt	ata	agt	
wt		tcc	tgg	ctg	ctg	ctg	ctc	ctg	ctt	tcc	ctc	tcc	tbc	ctc	caa	gcc	acg	gac	ttc	att	tct	
		L	*																			
env		tta	tga																			
wt		ctg	tga																			

FIGURE 6

06/717280

rTHY-1env

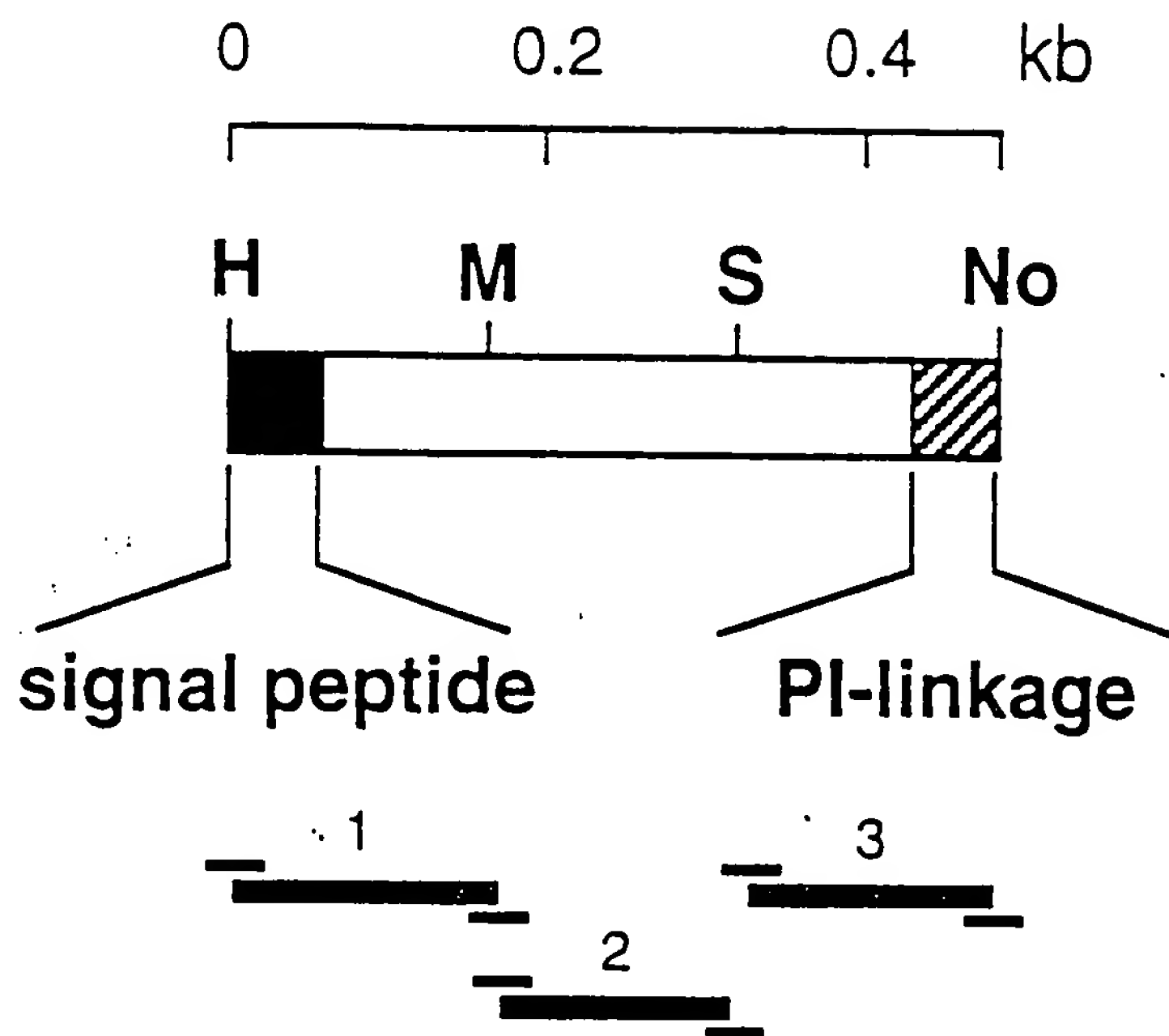


FIGURE 7

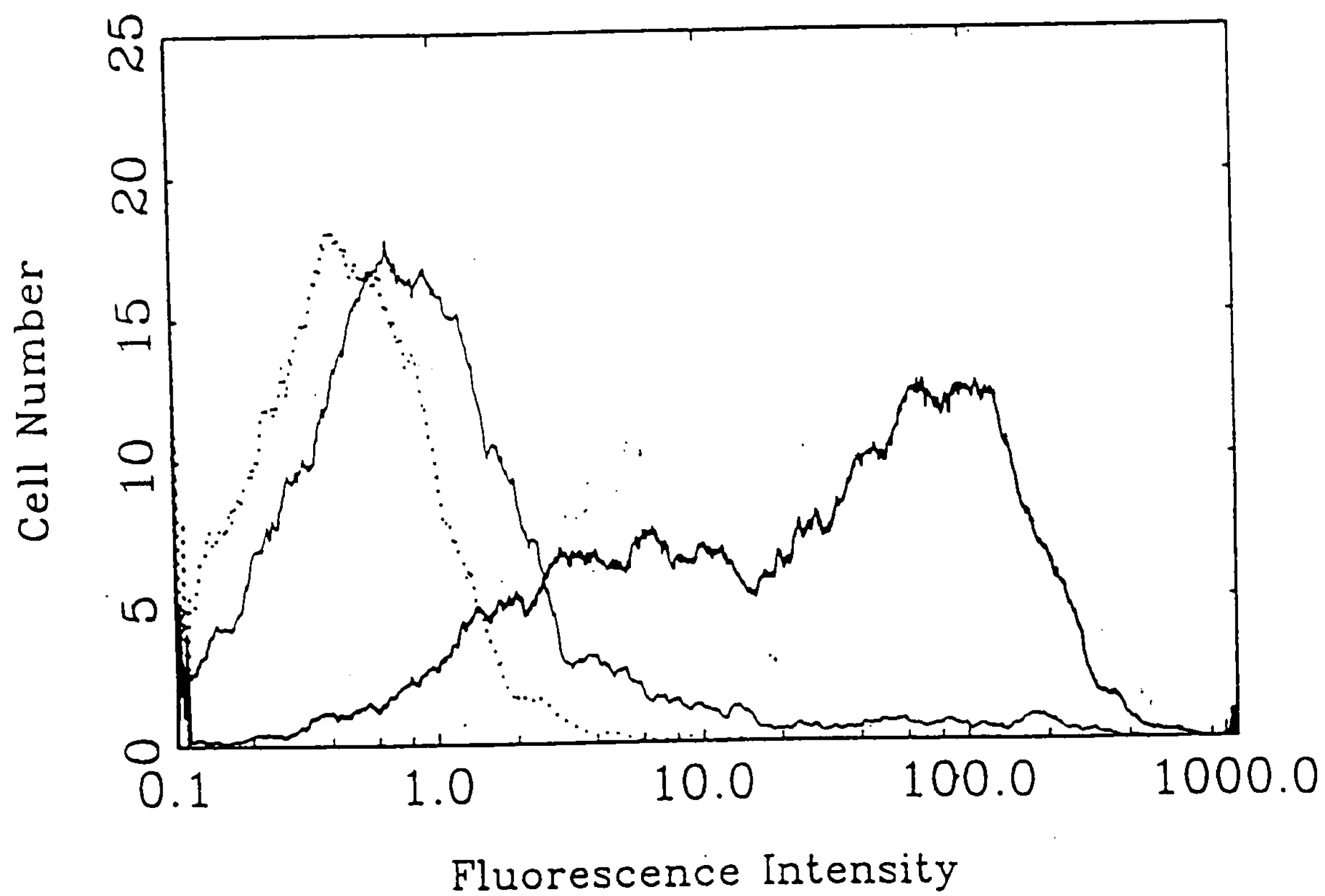


FIGURE 8

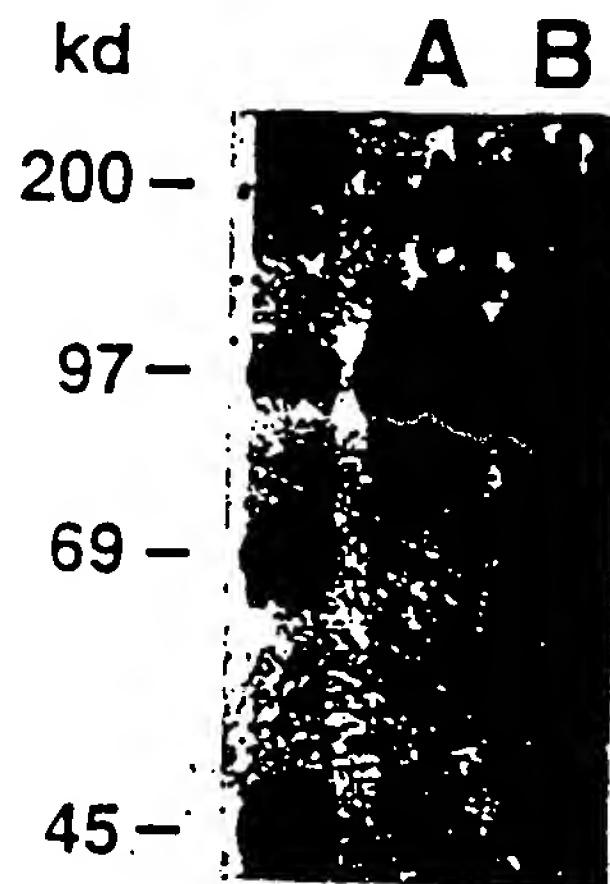
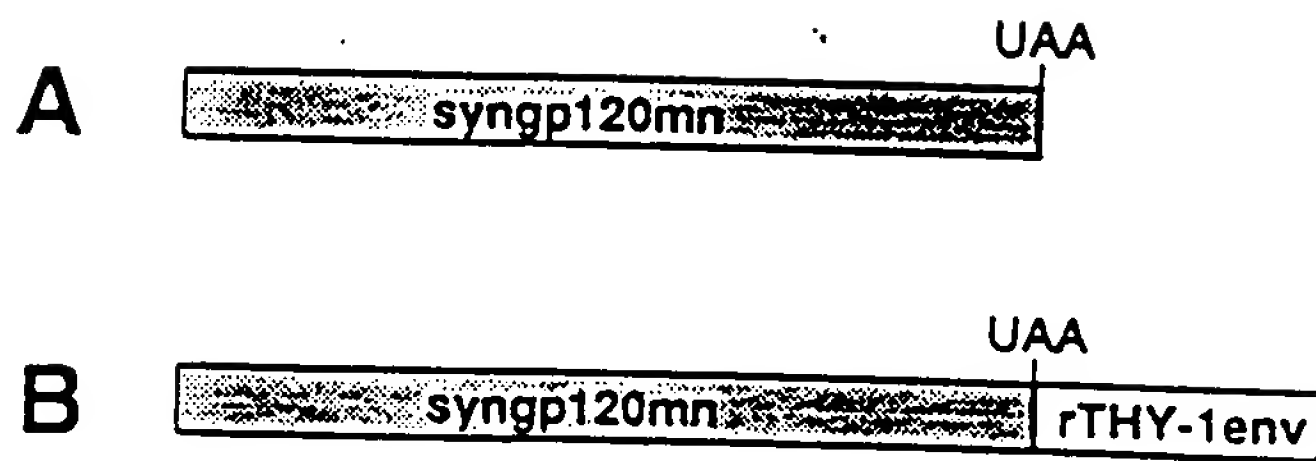
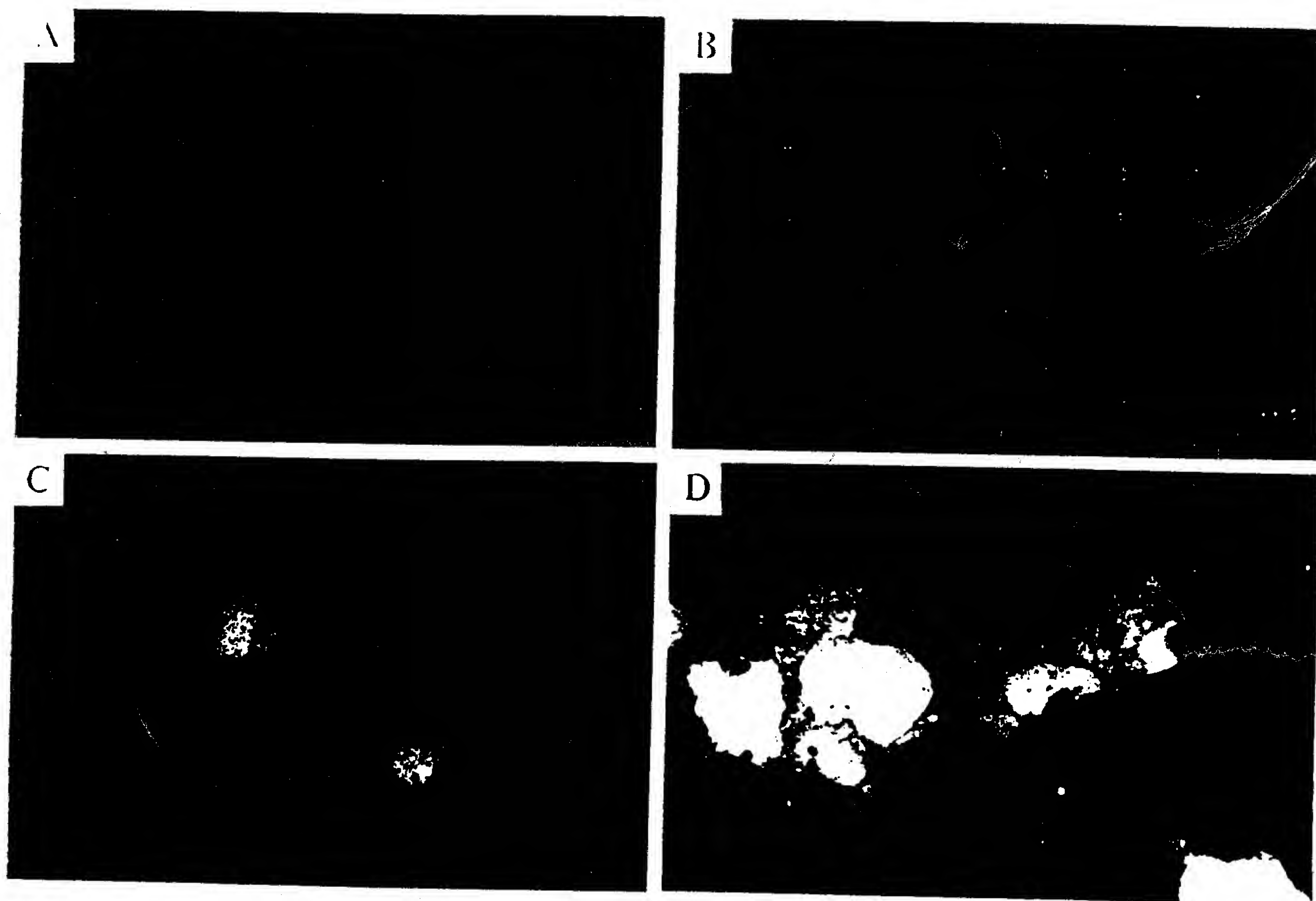
a*b*

FIGURE 9

FIG. 10



1 GAATTCACGC GTAAGCTTGC CGCCACCATG GTGAGCAAGG GCGAGGAGCT
51 GTTCACCGGG GTGGTGCCCA TCCTGGTCGA GCTGGACGGC GACGTGAACG
101 GCCACAAGTT CAGCGTGTCC GGCGAGGGCG AGGGCGATGC CACCTACGGC
151 AAGCTGACCC TGAAGTTCAT CTGCACCACC GGCAAGCTGC CCGTGCCCTG
201 GCCCACCCTC GTGACCACCT TCAGCTACGG CGTGCAGTGC TTCAGCCGCT
251 ACCCCGACCA CATGAAGCAG CACGACTTCT TCAAGTCCGC CATGCCCCGAA
301 GGCTACGTCC AGGAGCGCAC CATCTTCTTC AAGGACGACG GCAACTACAA
351 GACCCGCGCC GAGGTGAAGT TCGAGGGCGA CACCCTGGTG AACCGCATCG
401 AGCTGAAGGG CATCGACTTC AAGGAGGACG GCAACATCCT GGGGCACAAG
451 CTGGAGTACA ACTACAACAG CCACAACGTC TATATCATGG CCGACAAGCA
501 GAAGAACGGC ATCAAGGTGA ACTTCAAGAT CCGCCACAAC ATCGAGGACG
551 GCAGCGTGCA GCTCGCCGAC CACTACCAGC AGAACACCCC CATCGGCGAC
601 GGCCCCGTGC TGCTGCCCGA CAACCACTAC CTGAGCACCC AGTCCGCCCT
651 GAGCAAAGAC CCCAACGAGA AGCGCGATCA CATGGTCCTG CTGGAGTTCG
701 TGACCGCCGC CGGGATCACT CACGGCATGG ACGAGCTGTA CAAGTAAAGC
751 GGCCGCGGAT CC (SEQ ID NO: 40)

FIG. 11

Native Factor VIII B domain deleted gene segment inserted in the expression vector

1 AAGCTTAAAC CATGCCCATG GGGTCTCTGC AACCGCTGGC CACCTTGTAC
51 CTGCTGGGGA TGCTGGTCCG TTCCGTGCTA GCCGCCACCA GAAGATACTA
101 CCTGGGTGCA GTGGAAGTGT CATGGGACTA TATGCAAAGT GATCTCGGTG
151 AGCTGCCTGT GGACGCAAGA TTTCCTCCTA GAGTGCCAAA ATCTTTTCCA
201 TTCAACACCT CAGTCGTGTA CAAAAAGACT CTGTTTGTAG AATTCACGGA
251 TCACCTTTTC AACATCGCTA AGCCAAGGCC ACCCTGGATG GGTCTGCTAG
301 GTCCTACCAT CCAGGCTGAG GTTTATGATA CAGTGGTCAT TACACTTAAG
351 AACATGGCTT CCCATCCTGT CAGTCTTCAT GCTGTTGGTG TATCCTACTG
401 GAAAGCTTCT GAGGGAGCTG AATATGATGA TCAGACCAGT CAAAGGGAGA
451 AAGAAGATGA TAAAGTCTTC CCTGGTGGAA GCCATACATA TGTCTGGCAG
501 GTCCTGAAAG AGAATGGTCC AATGGCCTCT GACCCACTGT GCCTTACCTA
551 CTCATATCTT TCTCATGTGG ACCTGGTAAA AGACTTGAAT TCAGGCCTCA
601 TTGGAGCCCT ACTAGTATGT AGAGAAGGGA GTCTGGCCAA GGAAAAGACA
651 CAGACCTTGC ACAAATTTAT ACTACTTTTT GCTGTATTG ATGAAGGGAA
701 AAGTTGGCAC TCAGAAACAA AGAACTCCTT GATGCAGGAT AGGGATGCTG
751 CATCTGCTCG GGCCTGGCCT AAAATGCACA CAGTCAATGG TTATGTAAAC
801 AGGTCTCTGC CAGGTCTGAT TGGATGCCAC AGGAAATCAG TCTATTGGCA
851 TGTGATTGGA ATGGGCACCA CTCCTGAAGT GCACTCAATA TTCCTCGAAG
901 GTCACACATT TCTTGTGAGG AACCATCGCC AGGCGTCCTT GGAAATCTCG
951 CCAATAACTT TCCTTACTGC TCAAACACTC TTGATGGACC TTGGACAGTT
1001 TCTACTGTTT TGTCATATCT CTTCCCACCA ACATGATGGC ATGGAAGCTT
1051 ATGTCAAAGT AGACAGCTGT CCAGAGGAAC CCCAACTACG AATGAAAAAT
1101 AATGAAGAAG CGGAAGACTA TGATGATGAT CTTACTGATT CTGAAATGGA
1151 TGTGGTCAGG TTTGATGATG ACAACTCTCC TTCCTTTATC CAAATTGCTG
1201 CAGTTGCCAA GAAGCATCCT AAAACTTGGG TACATTACAT TGCTGCTGAA
1251 GAGGAGGACT GGGACTATGC TCCCTTAGTC CTCGCCCCCG ATGACAGAAG
1301 TTATAAAAGT CAATATTTGA ACAATGGCCC TCAGCGGATT GGTAGGAAGT
1351 ACAAAAAAGT CCGATTTATG GCATACACAG ATGAAACCTT TAAGACTCGT
1401 GAAGCTATTC AGCATGAATC AGGAATCTTG GGACCTTTAC TTTATGGGGA
1451 AGTTGGAGAC ACACTGTTGA TTATATTTAA GAATCAAGCA AGCAGACCAT
1501 ATAACATCTA CCCTCACGGA ATCACTGATG TCCGTCCTTT GTATTCAAGG
1551 AGATTACCAA AAGGTGTAAA ACATTTGAAG GATTTTCCAA TTCTGCCAGG
1601 AGAAATATTC AAATATAAAT GGACAGTGAC TGTAGAAGAT GGGCCAACTA
1651 AATCAGATCC TCGGTGCCTG ACCCGCTATT ACTCTAGTTT CGTTAATATG
1701 GAGAGAGATC TAGCTTCAGG ACTCATTGGC CCTCTCCTCA TCTGCTACAA
1751 AGAATCTGTA GATCAAAGAG GAAACCAGAT AATGTCAGAC AAGAGGAATG
1801 TCATCCTGTT TTCTGTATTT GATGAGAACC GAAGCTGGTA CCTCACAGAG
1851 AATATACAAC GCTTTCTCCC CAATCCAGCT GGAGTGCAGC TTGAGGATCC
1901 AGAGTTCCAA GCCTCCAACA TCATGCACAG CATCAATGGC TATGTTTTTG
1951 ATAGTTTGCA GTTGTCAGTT TGTTTGCATG AGGTGGCATA CTGGTACATT
2001 CTAAGCATTG GAGCACAGAC TGACTTCCTT TCTGTCTTCT TCTCTGGATA
2051 TACCTTCAA CACAAAATGG TCTATGAAGA CACACTCACC CTATTCCCAT
2101 TCTCAGGAGA AACTGTCTTC ATGTCGATGG AAAACCCAGG TCTATGGATT
2151 CTGGGGTGCC ACAACTCAGA CTTTCGGAAC AGAGGCATGA CCGCCTTACT
2201 GAAGGTTTCT AGTTGTGACA AGAACACTGG TGATTATTAC GAGGACAGTT
2251 ATGAAGATAT TTCAGCATAC TTGCTGAGTA AAAACAATGC CATTGAACCA
2301 AGAAGCTTCT CCCAGAATTC AAGACACCCT AGCACTAGGC AAAAGCAATT
2351 TAATGCCACC CCACCAGTCT TGAAACGCCA TCAACGGGAA ATAACTCGTA
2401 CTACTCTTCA GTCAGATCAA GAGGAAATTG ACTATGATGA TACCATATCA
2451 GTTGAAATGA AGAAGGAAGA TTTTGACATT TATGATGAGG ATGAAAATCA
2501 GAGCCCCCGC AGCTTTCAA AGAAAACACG ACACTATTTT ATTGCTGCAG
2551 TGGAGAGGCT CTGGGATTAT GGGATGAGTA GCTCCCCACA TGTCTAAGA
2601 AACAGGGCTC AGAGTGGCAG TGTCCCTCAG TTCAAGAAAG TTGTTTTCCA
2651 GGAATTTACT GATGGCTCCT TTAATCAGCC CTTATACCGT GGAGAACTAA
2701 ATGAACATTT GGGACTCCTG GGGCCATATA TAAGAGCAGA AGTTGAAGAT

Fig. 12

2751 AATATCATGG TAACTTTCAG AAATCAGGCC TCTCGTCCCT ATTCCTTCTA
 2801 TTCTAGCCTT ATTTCTTATG AGGAAGATCA GAGGCAAGGA GCAGAACCTA
 2851 GAAAAAACTT TGTCAAGCCT AATGAAACCA AAACCTTACTT TTGGAAAGTG
 2901 CAACATCATA TGGCACCCAC TAAAGATGAG TTTGACTGCA AAGCCTGGGC
 2951 TTATTTCTCT GATGTTGACC TGGAAAAAGA TGTGCACTCA GGCCTGATTG
 3001 GACCCCTTCT GGTCTGCCAC ACTAACACAC TGAACCCTGC TCATGGGAGA
 3051 CAAGTGACAG TACAGGAATT TGCTCTGTTT TTCACCATCT TTGATGAGAC
 3101 CAAAAGCTGG TACTTCACTG AAAATATGGA AAGAAACTGC AGGGCTCCCT
 3151 GCAATATCCA GATGGAAGAT CCCACTTTTA AAGAGAATTA TCGCTTCCAT
 3201 GCAATCAATG GCTACATAAT GGATACACTA CCTGGCTTAG TAATGGCTCA
 3251 GGATCAAAGG ATTCGATGGT ATCTGCTCAG CATGGGCAGC AATGAAAACA
 3301 TCCATTCTAT TCATTTTCTG GGACATGTGT TCACTGTACG AAAAAAAGAG
 3351 GAGTATAAAA TGGCACTGTA CAATCTCTAT CCAGGTGTTT TTGAGACAGT
 3401 GGAAATGTTA CCATCCAAAG CTGGAATTTG GCGGGTGGAA TGCCTTATTG
 3451 GCGAGCATCT ACATGCTGGG ATGAGCACAC TTTTCTGGT GTACAGCAAT
 3501 AAGTGTCAGA CTCCCCTGGG AATGGCTTCT GGACACATTA GAGATTTTCA
 3551 GATTACAGCT TCAGGACAAT ATGGACAGTG GGCCCCAAAG CTGGCCAGAC
 3601 TTCATTATTC CGGATCAATC AATGCCTGGA GCACCAAGGA GCCCTTTTCT
 3651 TGGATCAAGG TGGATCTGTT GGCACCAATG ATTATTCACG GCATCAAGAC
 3701 CCAGGGTGCC CGTCAGAAGT TCTCCAGCCT CTACATCTCT CAGTTTATCA
 3751 TCATGTATAG TCTTGATGGG AAGAAGTGGC AGACTTATCG AGGAAATTCC
 3801 ACTGGAACCT TAATGGTCTT CTTTGGCAAT GTGGATTCAT CTGGGATAAA
 3851 ACACAATATT TTTAACCCTC CAATTATTGC TCGATACATC CGTTTGCACC
 3901 CAACTCATTA TAGCATTTCG AGCACTCTTC GCATGGAGTT GATGGGCTGT
 3951 GATTTAAATA GTTGCAGCAT GCCATTGGGA ATGGAGAGTA AAGCAATATC
 4001 AGATGCACAG ATTACTGCTT CATCCTACTT TACCAATATG TTTGCCACCT
 4051 GGTCTCCTTC AAAAGCTCGA CTTACCTCC AAGGGAGGAG TAATGCCTGG
 4101 AGACCTCAGG TGAATAATCC AAAAGAGTGG CTGCAAGTGG ACTTCCAGAA
 4151 GACAATGAAA GTCACAGGAG TAACTACTCA GGGAGTAAAA TCTCTGCTTA
 4201 CCAGCATGTA TGTGAAGGAG TTCCTCATCT CCAGCAGTCA AGATGGCCAT
 4251 CAGTGGACTC TCTTTTTTCA GAATGGCAAA GTAAAGGTTT TTCAGGGAAA
 4301 TCAAGACTCC TTCACACCTG TGGTGAATC TCTAGACCCA CCGTTACTGA
 4351 CTCGCTACCT TCGAATTCAC CCCAGAGTT GGGTGCACCA GATTGCCCTG
 4401 AGGATGGAGG TTCTGGGCTG CGAGGCACAG GACCTCTACT GAGGGTGGCC
 4451 ACTGCAGCAC CTGCCACTGC CGTCACCTCT CCCTCCTCAG CTCCAGGGCA
 4501 GTGTCCCTCC CTGGCTTGCC TTCTACCTTT GTGCTAAATC CTAGCAGACA
 4551 CTGCCTTGAA GCCTCCTGAA TTAATATCA TCAGTCCTGC ATTTCTTTGG
 4601 TGGGGGGCCA GGAGGGTGCA TCCAATTTAA CTTAACTCTT ACCGTCGACC
 4651 TGCAGGCCCA ACGCGGCCGC

Fig. 12

(2 of 2)

Synthetic Factor VIII B domain deleted gene segment inserted in the expression vector

1 AAGCTTAAAC CATGCCCATG GGGTCTCTGC AACCGCTGGC CACCTTGATC
51 CTGCTGGGGA TGCTGGTCGC TTCCGTGCTA GCCGCCACCC GCCGCTACTA
101 CCTGGGCGCC GTGGAGCTGT CCTGGGACTA CATGCAGAGC GACCTGGGCG
151 AGCTCCCCGT GGACGCCCGC TTCCCCCCCC GCGTGCCCAA GAGCTTCCCC
201 TTCAACACCA GCGTGGTGTA CAAGAAAACC CTGTTCTGTG AGTTCACCGA
251 CCACCTGTTC AACATTGCCA AGCCGCGCCC CCCCTGGATG GGCCTGCTGG
301 GCCCCACCAT CCAGGCCGAG GTGTACGACA CCGTGGTGAT CACCCTGAAG
351 AACATGGCCA GCCACCCCGT CAGCCTGCAC GCCGTGGGCG TGAGCTACTG
401 GAAGGCCAGC GAGGGCGCCG AGTACGACGA CCAGACGTCC CAGCGCGAGA
451 AGGAGGACGA CAAGGTGTTC CCGGGGGGGA GCCACACCTA CGTGTGGCAG
501 GTGCTTAAGG AGAACGGCCC TATGGCCAGC GACCCCTGTG GCCTGACCTA
551 CAGCTACCTG AGCCACGTGG ACCTGGTGAA GGATCTGAAC AGCGGGCTGA
601 TCGGCGCCCT GCTGGTGTGT CGCGAGGGCA GCCTGGCCAA GGAGAAAACC
651 CAGACCCTGC ACAAGTTCAT CCTGCTGTTC GCCGTGTTCG ACGAGGGGAA
701 GAGCTGGCAC AGCGAGACTA AGAACAGCCT GATGCAGGAC CGCGACGCCG
751 CCAGCGCCCG CGCCTGGCCC AAGATGCACA CCGTTAACGG CTACGTGAAC
801 CGCAGCCTGC CCGGCCTGAT CGGCTGCCAC CGCAAGAGCG TGTACTGGCA
851 CGTCATCGGC ATGGGCACCA CCCCTGAGGT GCACAGCATC TTCCTGGAGG
901 GCCACACCTT CCTGGTGC GC AACCACCGCC AGGCCAGCCT GGAGATCAGC
951 CCCATCACCT TCCTGACTGC CCAGACCCTG CTGATGGACC TAGGCCAGTT
1001 CCTGCTGTTC TGCCACATCA GCAGCCACCA GCACGACGGC ATGGAGGCTT
1051 ACGTGAAGGT GGACAGCTGC CCCGAGGAGC CCCAGCTGCG CATGAAGAAC
1101 AACGAGGAGG CCGAGGACTA CGACGACGAC CTGACCGACA GCGAGATGGA
1151 TGTCGTACGC TTCGACGACG ACAACAGCCC CAGCTTCATC CAGATCCGCA
1201 GCGTGGCCAA GAAGCACCTT AAGACCTGGG TGCCTACATC CGCCGCCGAG
1251 GAGGAGGACT GGGACTACGC CCCGCTAGTA CTGGCCCCCG ACGACCGCAG
1301 CTACAAGAGC CAGTACCTGA ACAACGGCCC CCAGCGCATC GGCCGCAAGT
1351 ACAAGAAGGT GCGCTTCATG GCCTACACCG ACGAGACTTT CAAGACCCGC
1401 GAGGCCATCC AGCACGAGTC CGGCATCCTC GGCCCCCTGC TGTACGGCGA
1451 GGTGGGCGAC ACCCTGCTGA TCATCTTCAA GAACCAGGCC AGCAGGCCCT
1501 ACAACATCTA CCCCCACGGC ATCACCAGCG TGCGCCCCCT GTACAGCCGC
1551 CGCCTGCCCA AGGGCGTGAA GCACCTGAAG GACTTCCCCA TCCTGCCCGG
1601 CGAGATCTTC AAGTACAAGT GGACCGTGAC CGTGGAGGAC GGCCCCACCA
1651 AGAGCGACCC CCGCTGCCTG ACCCGCTACT ACAGCAGCTT CGTGAACATG
1701 GAGCGCGACC TGGCCTCCGG ACTGATCGGC CCCCTGCTGA TCTGCTACAA
1751 GGAGAGCGTG GACCAGCGCG GCAACCAGAT CATGAGCGAC AAGCGCAACG
1801 TGATCCTGTT CAGCGTGTTC GACGAGAACC GCAGCTGGTA TCTGACCGAG
1851 AACATCCAGC GCTTCCTGCC CAACCCCGCT GGCCTGCAGC TGGGAAGATCC
1901 CGAGTTCCAG GCCAGCAACA TCATGCACAG CATCAACGGC TACGTGTTCG
1951 ACAGCCTGCA GCTGAGCGTG TGCCTGCATG AGGTGGCCTA CTGGTACATC
2001 CTGAGCATCG GCGCCCAGAC CGACTTCCTG AGCGTGTTC TCTCCGGGTA
2051 TACCTTCAAG CACAAGATGG TGTACGAGGA CACCCTGACC CTGTTCCCCT
2101 TCTCCGGCGA GACTGTGTTC ATGTCTATGG AGAACCCCGG CCTGTGGATT
2151 CTGGGCTGCC ACAACAGCGA CTTCCGCAAC CGCGGCATGA CTGCCCTGCT
2201 GAAAGTCTCC AGCTGCGACA AGAACACCGG CGACTACTAC GAGGACAGCT
2251 ACGAGGACAT CTCCGCCTAC CTGCTGTCCA AGAACACGC CATCGAGCCC
2301 CGCTCCTTCT CCAAACCTC CCGCCACCCC AGCACGCGTC AGAAGCAGTT
2351 CAACGCCACC CCCCCGTGC TGAAGCGCCA CCAGCGCGAG ATCACCAGCA
2401 CCACCCTGCA AAGCGACCAG GAGGAGATCG ACTACGACGA CACCATCAGC
2451 GTGGAGATGA AGAAGGAGGA CTTGACATC TACGACGAGG ACGAGAACCA
2501 GAGCCCCCGC TCCTTCCAAA AGAAAACCG CCACTACTTC ATCGCCGCCG
2551 TGGAGCGCCT GTGGGACTAC GGCATGAGCA GCAGCCCCCA CGTCTGCGC
2601 AACCGCGCCC AGAGCGGCAG CGTGCCCCAG TTCAAGAAGG TGGTGTTCCT
2651 GGAGTTCACC GACGGCAGCT TCACCCAGCC CCTGTACCGC GGCGAGCTGA
2701 ACGAGCACCT GGGCCTGCTC GGCCCCTACA TCCGCGCCGA GGTGGAGGAC

Fig. 13

2751 AACATCATGG TGACCTTCCG CAACCAAGCC TCCCGGCCCT ACTCCTTCTA
2801 CTCCTCCCTG ATCAGCTACG AGGAGGACCA GCGCCAGGGC GCCGAGCCCC
2851 GCAAGAACTT CGTGAAGCCC AACGAGACTA AGACCTACTT CTGGAAGGTG
2901 CAGCACCACA TGGCCCCCAC CAAGGACGAG TTCGACTGCA AGGCCTGGGC
2951 CTACTTCAGC GACGTGGACC TGGAGAAGGA CGTGACACAGC GGCCTGATCG
3001 GCCCCCTGCT GGTGTGCCAC ACCAACACCC TGAACCCCCC CCACGGGAGG
3051 CAGGTGACTG TGCAGGAATT TGCCCTGTTC TTCACCATCT TCGACGAGAC
3101 TAAGAGCTGG TACTTCACCG AGAACATGGA GCGCAACTGC CGCGCCCCCT
3151 GCAACATCCA GATGGAAGAT CCCACCTTCA AGGAGAACTA CCGCTTCCAC
3201 GCCATCAACG GCTACATCAT GGACACCCTG CCCGGCCTGG TGATGGCCCA
3251 GGACCAGCGC ATCCGCTGGT ACCTGCTGTC TATGGGCAGC AACGAGAACA
3301 TCCACAGCAT CCACTTCAGC GGCCACGTTT TCACCGTGCG CAAGAAGGAG
3351 GAGTACAAGA TGGCCCTGTA CAACCTGTAC CCCGGCGTGT TCGAGACTGT
3401 GGAGATGCTG CCCAGCAAGG CCGGGATCTG GCGCGTGGAG TGCCTGATCG
3451 GCGAGCACCT GCACGCCGGC ATGAGCACCC TGTTCTTGGT GTACAGCAAC
3501 AAGTGCCAGA CCCCCCTGGG CATGGCCAGC GGCCACATCC GCGACTTCCA
3551 GATCACCGCC AGCGGCCAGT ACGGCCAGTG GGCTCCCAAG CTGGCCCCGCC
3601 TGCATAACAG CGGCAGCATC AACGCCTGGT CGACCAAGGA GCCCTTCTCC
3651 TGGATCAAGG TGGACCTGCT GGCCCCCATG ATCATCCACG GCATCAAGAC
3701 CCAGGGCGCC CGCCAGAAGT TCAGCAGCCT GTACATCAGC CAGTTCATCA
3751 TCATGTACTC TCTAGACGGC AAGAAGTGGC AGACCTACCG CGGCAACAGC
3801 ACCGGCACCC TGATGGTGTT CTTCGGCAAC GTGGACAGCA GCGGCATCAA
3851 GCACAAACATC TTCAACCCCC CCATCATCGC CCGCTACATC CGCCTGCACC
3901 CCACCCACTA CAGCATCCGC AGCACCTGCG GCATGGAGCT GATGGGCTGC
3951 GACCTGAACA GCTGCAGCAT GCCCCTGGGC ATGGAGAGCA AGGCCATCAG
4001 CGACGCCCAG ATCACCGCCT CCAGCTACTT CACCAACATG TTCGCCACCT
4051 GGAGCCCCAG CAAGGCCCGC CTGCACCTGC AGGGCCGCGC CAACGCCTGG
4101 CGCCCCCAGG TGAACAACCC CAAGGAGTGG CTGCAGGTGG ACTTCCAGAA
4151 AACCATGAAG GTGACTGGCG TGACCACCCA GGGCGTCAAG AGCCTGCTGA
4201 CCAGCATGTA CGTGAAGGAG TTCCTGATCA GCAGCAGCCA GGACGGCCAC
4251 CAGTGGACCC TGTTCTTCCA AAACGGCAAG GTGAAGGTGT TCCAGGGCAA
4301 CCAGGACAGC TTCACACCGG TCGTGAACAG CCTGGACCCC CCCCTGCTGA
4351 CCCGCTACCT GCGCATCCAC CCCAGAGCT GGGTGCACCA GATCGCCCTG
4401 CGCATGGAGG TGCTGGGCTG CGAGGCCCAG GACCTGTACT GAAGCGGCCG
4451 C

Fig. 13